# **REMARKS**

The present application relates to inbred maize line PH77V. Claims 17-20, 33, 36, and 51-53 have been canceled. Claims 9, 11, 13, 14, 21-22, 28, 41, 42, 47-49, and 50 have been amended. New claims 54-72 have been added. No new matter has been added by the present amendment. Applicant respectfully requests consideration of the following remarks.

## **Detailed Action**

## A. Claim and Specification Objections

Applicant acknowledges the objection to claims 8 and 27 as withdrawn in light of the claim amendments. Applicant further acknowledges the rejections of claims 1-49 under 35 U.S.C. § 112, second paragraph, as withdrawn in light of the claim amendments. The rejection of claims 1-49 under 35 U.S.C. § 112, first paragraph, are acknowledged as withdrawn in light of its deposit. Applicant acknowledges the rejection of claims 18-20 and 47-49 under 35 U.S.C. § 112, first paragraph, for lack of enablement, as withdrawn.

# Rejections under 35 U.S.C. § 112, Second Paragraph

Claims 22, 33, 47-49, and 50-53 stand rejected under 35 U.S.C. § 112, second paragraph, as being indefinite for failing to particularly point out and distinctly claim the subject matter which Applicant regards as the invention.

Claim 22 is indefinite for the recitation "said plant has essentially the same morphology and physiology of inbred maize line PH77V other than the trait of male sterility." The Examiner states it is unclear what is meant by "essentially the same".

Applicant has amended claim 22 to delete the terminology "essentially the same morphology and physiology of inbred maize line PH77V", thus alleviating this rejection. Applicant further asserts that previously submitted Attachment C was presented as evidence of what those of ordinary skill in the art accept as the definition of "essentially derived variety" not what UPOV allows.

Claim 33 stands rejected as indefinite for the recitation "pedigree of said PH77V-progeny maize plant is within 2 or less crosses".

Applicant has now canceled claim 33, thereby alleviating this rejection.

The Examiner rejects claim 47 for the recitation "essentially unchanged" as being indefinite.

Applicant has amended claim 47 to delete the terminology "essentially unchanged", alleviating this rejection.

Claim 50 is indefinite as the Examiner states it broadens the scope of the claims from which it depends.

Applicant traverses this rejection. Applicant asserts claim 50 does not broaden the claim from which it depends but rather narrows the claim to further teach a maize plant, or parts thereof, produced by growing the seed of claim 1 further comprising a gene that confers a specific trait not previously exhibited in said plant. Thus Applicants request reconsideration and withdrawal of the rejection.

The Examiner rejects claims 51-53 for the recitation "inbred PH77V maize plant conferring a backcrossed trait of claim 51" as indefinite. The Examiner states it is not clear what is meant by a plant that confers a backcrossed trait.

Applicant has canceled claims 51-53, alleviating this rejection.

Claim 52 is rejected for attempting to limit the "inbred PH77V maize plant conferring a backcrossed trait of claim 51" when claim 51 is directed to a method. Claim 52 is also indefinite for the recitation "essentially the same traits".

Applicant has canceled claim 52, thus alleviating these rejections.

In light of the above amendments and remarks, Applicant respectfully requests reconsideration and withdrawal of the rejections under 35 U.S.C. § 112, second paragraph.

## Rejections Under 35 U.S.C. § 112, First Paragraph

Claims 9-20, 28, 29, 33-39, 41-43, and 47-49 remain rejected and claims 22 and 50-53 stands rejected under 35 U.S.C. § 112, first paragraph, as containing subject matter which was not described in the specification in such a way as to reasonably convey to one skilled in the relevant art that the inventor(s), at the time the application was filed, had possession of the claimed invention, for the reasons of record stated in the Office Action mailed September 4, 2002, under item 4.

The Applicant traverses the rejection. Claims 17-20, 33, 36, and 51-53 have been canceled. Claims 9, 11, 13, 14, 21-22, 28, 41, 42, 47-49, and 50 have been amended. New claims 54-72 have been added.

The Examiner rejects claims 9, 10, 28, and 29, that claim the F1 hybrid seed and F1 hybrid plant made with PH77V as a parent. Claims 9 and 28 have been amended and claims 10 and 29 refer directly to claims 9 and 28. One of ordinary skill in the art would know how to cross PH77V with another maize plant. Applicant asserts it is well understood by one skilled in the art that maize is a diploid plant species thereby comprising two sets of chromosomes. The F1 hybrid seed and plant produced using PH77V, regardless of the other maize plant used, is identifiable because it will have a single set of individual maize chromosomes coming from PH77V. Therefore, it would be clear to one ordinarily skilled in the art that the addition of claim terminology --having a single set of maize chromosomes from PH77V-- in claims 9 and 28 would certainly be understood. In addition, one of ordinary skill in the art would be able to run a molecular profile on PH77V and the F1 hybrid and be able to identify the F1 hybrid as being produced from PH77V. PH77V is a homozygous inbred plant. When the ovule or pollen is generated from this plant, it will be haploid and will contain one complete set of chromosomes from PH77V. Upon fertilization, the resulting zygote will receive one set of chromosomes from the parent inbred plant resulting in the diploid zygote. Inbred PH77V has a unique set of genes present on its chromosomes and this unique set is also present in the hybrid.

As stated in the specification on page 16, lines 8-34, there are many laboratory-based techniques available for the analysis comparison and characterization of plant genotype such as Restriction Length Polymorphisms (RFLPs) and Simple Sequence Repeats (SSRs). Such techniques may be used to identify whether or not PH77V was used to develop a hybrid. Any person of skill in the art could run a molecular profile of PH77V based upon the deposit Applicant has made. Therefore, it would be routine to one of ordinary skill in the art to run the profile of a hybrid plant and determine whether or not PH77V was used as a parent.

Claim 17 has also been canceled. Claim 14 has been amended providing one of skilled in the art sufficient description to evaluate the presence of the claimed traits.

Claims 15 and 16 remain pending and are to methods of making a maize plant through the utilization of PH77V. Applicant points out that anyone of skill in the art would know how to utilize the well established breeding methods with PH77V. Description of such occurs

throughout the specification and descriptions can also be found in introductory plant breeding books. As stated in the written description guidelines, an old process performed with a novel material is novel in and of itself. 66 Federal Register 1099, Vol. 66, No. 4 (January 5, 2001). Further, claims 15 and 16 are methods patentable pursuant to the written description guidelines. See Example 10, Revised Interim Written Description Guidelines Training Materials, in which claim 1 therein is indicated as allowable.

Claim 40 is allowed. Claim 40 is to the method of producing a first generation (F1) PH77V-progeny maize plant. Claim 41 is to the first generation (F1) PH77V-progeny maize plant produced by the method of claim 40. The first generation (F1), or hybrid, is identifiable through both breeding records and molecular marker techniques as discussed above. Further as described herein, claim 41 requires that the hybrid have the complete set of PH77V maize chromosomes which are present in duplicate form in the inbred parent. Claim 42 is to the method of selfing the first generation (F1) PH77V for successive filial generations. Claim 43 is to the PH77V inbred progeny maize plant produced by the method of claim 42. Applicant asserts claim 42 is a method patentable pursuant to the written description guidelines. Further, claim 43 is written in the proper product-by-process format as taught in the written description guidelines. See Example 10, Revised Interim Written Description Guidelines Training Materials. This is a basic and well known breeding methodology, and the use of this methodology with PH77V is described in the specification on page 21, lines 16 to 31.

As stated in Openshaw *et al.* submitted herewith, "[t]he backcross breeding procedure has been used widely to transfer simply inherited traits into elite genotypes. ... Today, backcrossing is being used to transfer genes introduced by such techniques as transformation or mutation into appropriate germplasm." Openshaw *et al.* further notes breeders, by using molecular markers, may obtain a very high degree of genome identity between the backcross conversion and the recurrent parent after two backcrosses. See Marker-assisted Selection in Backcross Breeding, Openshaw, S.J. et al. Marker-assisted selection in backcross breeding. In: Proceedings Symposium of the Analysis of Molecular Data, August 1994, pages 41-43. Crop Science Society of America, Corvallis, OR (1994) included as <u>Appendix A</u>. The backcross method has been successfully used since the 1950's (see pages 585-586 of Wych, 1988 included in the Information Disclosure Statement). Thus Applicant asserts such mutant genes or transgenes may be introgressed into elite lines such as PH77V without undue experimentation. As further

evidence of this, Poehlman *et al.* (1995) on page 334, submitted in the Information Disclosure Statement, states that, "[a] backcross-derived inbred line fits into the same hybrid combination as the recurrent parent inbred line and contributes the effect of the additional gene added through the backcross." In addition, Wych (1988) on page 585-86, discusses how the male sterility trait is routinely backcrossed into an inbred line and how this is used to produce a sterile/fertile blend of an F1 hybrid in order to reduce seed production costs. In fact, many commercial products are produced in this manner, and those of ordinary skill in the art consider the F1 hybrid produced with the male sterile (backcross conversion) inbred to be the same variety as the F1 hybrid produced with the non-backcross conversion inbred.

The Examiner states that "the molecular profile of PH77V is not described in the specification." Applicant respectfully traverses this rejection. As described in the specification, lines 8-34, on page 16, the seed deposit allows one of ordinary skill to run a molecular profile of PH77V. Thus, one of ordinary skill in the art may test material they desire to use in breeding to determine if it is PH77V. An SSR profile is an inherent feature of inbred line PH77V, a representative sample of which has been deposited with the ATCC. For example, see Ex parte Marsili, Rosetti, and Pasqualucci, 214 USPQ 904 (1972), in which The Board, relying on well established cases of In re Nathan et al., 51 CCPA 1059, 328 F.2d 1005, 140 USPQ 601 (1964); In re Sulkowski, 487 F.2d 920, 180 USPQ 46 (CCPA 1973); Spero v. Ringold, 54 CCPA 1407, 377 F.2d. 652, 153 USPQ 726 (1967), and Petisi et al. v. Rennhard et al., 53 CCPA 1452, 363 F. 2d 903, 150 USPQ 669 (1966), concluded that the "products described, exemplified and claimed by Appellants inherently had and have now the structure given in the amendment in question". Applicant is willing to provide the molecular marker profile of PH77V.

The Examiner states that, "describing a plant that by saying it expresses 2 particular traits does not distinguish it from any other plant that expresses the same traits." Applicant points out that those claims referenced by the Examiner require the utilization of PH77V to develop such plant. However, in order to expedite prosecution the claims identifying progeny by phenotypic traits have been amended to further define and clearly claim the ancestor of PH77V expressing all the claimed traits, thereby alleviating the rejection.

The Examiner also states that the morphological and physiological traits of PH77V progeny are not described. The test of written description does not require a morphological and physiological description. Rather, it is whether subject matter was described in such a way to

convey to one of ordinary skill in the art that the inventor had possession of the claimed invention. While PVP is distinct from patents, the scope of protection conferred by PVP provides a clear indication that breeders of ordinary skill in the art consider F1 hybrids, backcross conversions and transgenic conversions to be within the scope of the invention of the variety itself. See previously submitted <a href="Attachment C">Attachment C</a>. Applicant asserts that Attachment C was presented as evidence of what those of ordinary skill in the art accept as the definition of "essentially derived variety" not what UPOV permits. These derivatives, variants and closely related progeny easily and routinely created through the use of this newly developed line are encompassed within the scope of the invention of the variety itself. The fact that the progeny have not been created does not prevent them from being protected in this manner. As stated in MPEP § 2163(3)(a), "An invention may be complete and ready for patenting before it has actually been reduced to practice."

The Examiner also rejects claims 37-39 under 35 U.S.C § 112, first paragraph. Claims 37-39 are directed to growing out an F1 hybrid in which PH77V is a parent and searching for PH77V inbred seed. Due to the imperfect process of seed production, parent seed can sometimes be contained in the hybrid seed bag. This claim covers the method of searching for inbred PH77V seed within a bag of hybrid seed. The method is clearly described in the specification on page 5, line 21 through line 7 on page 6. One of ordinary skill in the art can practice such a method without undue experimentation. The Applicant requests that the Examiner withdraw his rejection to claims 37-39.

The Examiner rejects claims to transgenic PH77V plants and PH77V plants comprising single gene conversions. New claims 54-72 are drawn to methods and to the products produced by those methods. The product by process claims are further limited by specified traits conferred by mutant genes or transgenes, which include the traits of insect resistance, herbicide resistance, disease resistance, and male sterility.

Applicant respectfully points out that examples of transgenes, mutant genes, genes, and traits that can be introduced into the PH77V are given in the application on page 21, lines 16-34, and also on page 22, line 34, through page 35, line 2. The Examiner suggests that the claims be amended to include a list of transgenes. In order to expedite prosecution new claims 54-72 list the type of traits that may be conferred. However it should be noted that PH77V comprising a mutant gene or a transgene, even if it is for a transcription factor, is distinct from another inbred

line comprising that same mutant gene or transgene and still retains the benefit of Applicant's invention.

In light of the above amendments and remarks, Applicant respectfully requests reconsideration and withdrawal of the rejections to claims 9-20, 22, 28, 29, 33-39, 41-43, 47-49, and 50-53 under 35 U.S.C. § 112, first paragraph.

## <u>Issues Under 35 U.S.C. § 102/103</u>

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Claims 9, 10, 13, 14, 17, 22, 28, 29, 33, 36, 41-43, 47-49 remain and claim 52 stand rejected under 35 U.S.C. § 102(b) as anticipated by or, in the alternative, under 35 U.S.C. § 103(a) as obvious over Puskaric (U.S. Patent No. 5,977,456), for the reasons of record stated in the Office Action mailed September 4, 2002, under item 7.

Applicant respectfully traverses this rejection. Applicant has canceled claims 17, 33, 36, and 44-46, thereby alleviating this rejection. Further, Applicant has amended claims 9, 13, 14, 22, 28, 41, 42, and 47-49 that are to the first generation (F1) developed from crossing PH77V with a second plant. Further claim 43 is a method to the PH77V inbred progeny maize plants produced by the method of claim 42. As explained earlier, the hybrid developed from inbred PH77V as a parent retain the same unique assemblage of genetic material present in duplicate form in the inbred. This contributes predictable traits to the hybrid as described in the specification.

The Examiner states that product-by-process claims may be properly rejected over prior art teaching the same product produced by a different process." It is erroneous to assume that PH1M7 is the same as PH77V either phenotypically or genetically, and Applicant has disclosed information which may be used to distinguish this both from phenotype and genetic profile. PH1M7 is not PH77V, nor can PH1M7 be created through the use of PH77V with one breeding cross. Thus, claim 41 and 43 are not anticipated by PH1M7. Further, Applicant submits In re Thorpe, states that "a product by process claim may be properly rejected over prior art teaching the same product produced by a different process", as noted by the Examiner. In re Thorpe, 227 USPQ. 964, 966 (Fed. Cir. 1985). However, Applicant submits that this is not the same product physiologically or morphologically as the cited prior art as can be evidenced by one skilled in the art through analysis of the data tables in each and the differences described in the Amendment of January 6, 2003. In addition, it is impermissible to use hindsight reconstruction and the benefit

of Applicant's disclosure to pick among pieces which are present in the art, there must be some suggestion to make the combination and an expectation of success. In re Vaeck, 20 USPQ2d 1434 (Fed. Cir. 1991). Moreover, Applicant claims a method of making a plant which did not previously exist. Pursuant to the recent Federal Circuit decision, Elan Pharmaceuticals, Inc. v. Mayo Foundation for Medical Education & Research, 304 F.3d 1221, (Fed. Cir. 2002), "a novel patented product is not "anticipated" if it did not previously exist." Id. This is the case whether or not the process for making the new product is generally known. Id. The invention PH77V has not previously existed therefore Applicant strongly asserts that neither the suggestion of the claimed unique invention of the present application nor the expectation of success is taught for one ordinarily skilled in the art in the reference cited by the Examiner.

In light of the above, Applicant respectfully requests that the Examiner reconsider and withdraw the rejections to claims 9, 10, 13, 14, 17, 22, 28, 29, 33, 36, 41-43, 47-49, and 52 under 35 U.S.C. § 102(b) as anticipated by or, in the alternative, under 35 U.S.C. §103(a) as obvious over Puskaric (U.S. Patent No. 5,977,456).

# **Summary**

Applicant acknowledges that claims 1-8, 21, 23-27, and 40 are allowed.

# Conclusion

In conclusion, Applicant submits in light of the above amendments and remarks, the claims as amended are in a condition for allowance, and reconsideration is respectfully requested. If it is felt that it would aid in prosecution, the Examiner is invited to contact the undersigned at the number indicated to discuss any outstanding issues.

No fees or extensions of time are believed to be due in connection with this amendment; however, consider this a request for any extension inadvertently omitted, and charge any additional fees to Deposit Account No. 26-0084.

Reconsideration and allowance is respectfully requested.

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Respectfully submitted,

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# Marker-assisted Selection in Backcross Breeding

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Abstract. The backcross breeding procedure has been used widely to transfer simply inherited traits into elite genotypes. Genetic markers can increase the effectiveness of backcrossing by 1) increasing the probability of obtaining a suitable conversion, and 2) decreasing the time required to achieve an acceptable recovery. Simulation and field results indicated that, for a genome consisting of ten 200-cM chromosomes, basing selection on 40 or 80 markers in 50 BC individuals that carry the allele being transferred can reduce the number of backcross generations needed from about seven to three.

to transfer simply inherited traits into elite genotypes. Usually, the trait being transferred is controlled by a single gene, but highly heritable traits that are more complexly inherited have also been transferred successfully by backerossing; for example, mainrity in maize (Rinke and Sentz, 1961; Shaver, 1976). Today, backerossing is being used to transfer genes introduced by such techniques as transformation or mutation into appropriate germplasm.

Several plant breeding textbooks give good descriptions of the backcross procedure (Allard, 1960; Fehr, 1987). A donor parent (DP) carrying a trait of interest is crossed to the recurrent parent (RP), an elite line that is lacking the trait. The F, is crossed back to the RP to produce the BC, generation. In the BC, and subsequent backcross generations, selected individuals carrying the gene being transferred are backcrossed to the RP. The expected proportion of DP genome is reduced by half with each generation of backcrossing. Ignoring effects of linkage to the selected DP allele being transferred, the percentage recurrent parent (%RP) genome expected in each backcross generation is calculated as:

 $%RP = 100[1 - (0.5)^{n+1}]$ 

where n is the number of backcrosses.

Backcrossing of selected plants to the RP can be repeated each cycle until a line is obtained that is essentially a version of the RP that includes the introgressed allele. After six backcrosses, the expected recovery is >99% (Table 1).

Until recently, discussions of the recovery of the RP genome during backcrossing have emphasized the expected values for %RP shown in Table 1, and have largely ignored the genetic variation for %RP that exists around the expected mean. With the development of genetic markers capable of providing good genome coverage, there has been interest in taking advantage of that variation to increase the efficiency of backcrossing.

Selection for RP marker alleles can increase greatly the effectiveness of backcross programs by allowing the breeder to 1) select backcross plants that have a higher proportion of RP genome, and 2) select backcross individuals that are better conversions near a mapped donor allele being transferred (i.e., select for less linkage drag). Expressed in practical terms, using genetic markers to assist backcrossing can 1) increase the probability of obtaining a suitable conversion, and 2) decrease the time required to achieve an acceptable recovery.

Issues to consider when planning a marker-assisted backcross program include 1) the time advantage of using markers to assist backcrossing, 2) the number of markers needed, and 3) the number of genotypes to evaluate. In this report, we use results from previous literature, computer simulation, and empirical studies to provide some guidelines.

Table 1. Expected recovery of recurrent parent (RP) genome during backcrossing, assuming no linkage to the gene being transferred.

Generation	% RP		
F,	50.0000		
BC,	75.0000		
	87.5000		
BC,	93.7500		
BC,	96,8750		
BC, BC,	98.4375		
ac,	99.2188 '		
BC, .	99,6094		

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Analysis of Molecular Marker Data

#### Materials and methods

The maize genome was the model for the simulation. The simulated genome contained ten 200-cM chromosomes. Simulation of crossing over was based on a Poisson distribution with a mean of 2.0 ( $\lambda$  = 2) (Hanson, 1959), which, on average, generated one cross over for every 100-cM length. The simulations reported here assume no interference. Codominant genetic markers were evenly distributed in the genome and sites of the donor gene were randomly assigned to genome locations.

Simulations were conducted with the following parameters:

Number of progeny: 100 or 500.

Backcross generations: BC<sub>1</sub>, BC<sub>2</sub>, and BC<sub>3</sub>. Number of markers: 20, 40, 80, or 100.

Number selected to form the next BC generation: 1 or 5.

Selection was based on 1) presence of the donor allele and 2) high %RP). %RP was calculated as the average of the (one or five) selected individuals. Values presented are the mean of 50 simulations.

#### Results

In the computer simulation study, all methods modeled greatly increased the speed of recovering the RP genome compared to the expected recovery with no marker-assisted selection (compare Tables 1 and 2). At least 80 markers were required to recover 99% of the RP genome in just three BC generations (Table 2). Use of at least 80 markers and 500 progeny allowed recovery of 98% RP in just two BC generations. Response to selection was diminished only slightly by spreading the effort over five selections. Using markers, the number of backcross generations needed to convert an inbred is

reduced from about seven to three.

By the BC<sub>1</sub> generation, there appears to be no practical advantage to using 500 vs. 100 individuals. If the presence of the donor trait in the backcross individuals can be ascertained before markers are genotyped, then only half the number of individuals indicated in the tables will need to be analyzed.

When a small number of markers are used, they quickly became non-informative; i.e., selection causes the marker loci to became fixed for the RP type before the rest of the genome is fully converted (Table 3; Hospital et al., 1992). This situation was most prominent in the larger populations, where a higher selection intensity placed more selection pressure upon the marker loci. Accordingly, it is of interest to consider how closely the estimation of %RP based on markers reflects the actual genome composition. The combination of estimation of %RP based on fewer markers and subsequent selection tends to bias the estimates upward (compare Tables 2 and 3).

The results from the simulation compare well with real field data. In a typical example, 50 BC, plants carrying the gene being transferred were genotyped at 83 polymorphic RFLP loci (note that this corresponds to a population size of 100 unselected plants in Tables 2 and 3). The five best BC, recoveries had estimated %RP values of 85.9%, 82.7%, 82.0%, 81.4%, and 81.2%. After evaluating 10 BC, plants from each selected BC, the best BC, recovery had an estimated %RP of 94.6%.

#### Discussion

The simulations (Table 2; Hospital et al., 1992) and our experience indicate that four markers per 200-cM chromosome is adequate to greatly increase the effectiveness of selection in the BC<sub>1</sub>. However, using only four markers per 200 cM will likely make it very difficult to map the location of the gene of interest. Adequate summarization of the data is an important

Table 2. Percent recurrent parent genome during marker-assisted backernssing.

Generation	100 Progeny No. markers					500 Progeny			
					No. markers				
	20	40	80	100	20	40	20	100	
		. —	Or	se selected	•				
BC,	84.5	84.5	84.2	88.0	89.9	90.7	90.2	90.5	
BC,	95.0	95.2	. 95.8	97.2	96.5	97.7	98.5	98.6	
BC, BC, BC,	97.4	97.6	98.9	99.2	97.7	98.3	99.4	99.5	
			Fin	ve selected			•		
BC,	82.9	85.1	84.9	84.7	87.7	88.1	88.9	88.9	
BC,	93.7	95.0	95.8	95.7	95.5	96.8	97.8	97.9	
BC, BC, BC,	97.1	98.3	98.8	98.9	97.3	98.5	99.3	99,3	

Table 3. Estimates of percent recurrent parent genome, based on marker loci.

Generation	No. markers				500 Progeny No. markers			
				Or	e selected			
BC,	98.7	97.8	95.6	97.2	100.0	99.1	98.6	98.0
BC, BC,	100.0	99.8	99.3	99.5	100.0	100.0	99.9	98.2
			Fit	ve selected.				
BC,	96.4	96.5	96.2	95.8	100.0	98.5	98.3	98.2
BC, BC,	99.9	99.8	99.3	99.1	100.0	100.0	99.9	99.8

igan of a marker-assisted backcross program. Ideally, the markers used can supply data that can be represented as alleles of loci with known map position. Estimation of %RP, mapping the position of the locus of interest, and graphical display of the results (Young and Tanksley, 1989) are all useful in underganding and controlling the specific backcross experiment being conducted.

It appears that, with the use of genetic markers, the portion of the RP genome that is not linked to the allele being transferred can be recovered quickly and with confidence. The recovery of RP will be slower on the chromosome carrying the gene of interest. A considerable amount of linkage drag is expected to accompany selection for the DP allele in a backgross program. For a locus located in the middle of a 200-cM chromosome, the length of the DP chromosome segment accompanying selection is expected to be 126, 63, and 28 cM in the BC, BC, and BC, generations, respectively (Hanson, 1959; Naveira and Barbadilla, 1992). Our observations support the recommendation of Hospital et al. (1992) that preference be given to the selection for recombinants proximal to the allele of interest, but that selection for recovery of the RP elsewhere in the genome also be considered. This two-stage selection can probably be done quite effectively ad hoc by the breeder once the data is adequately summarized; however, Hospital et al.

suggest ways to incorporate the two criteria into a selection index such that each component of selection is assured appropriate weighting.

Use of genetic markers can greatly increase the effectiveness of backcrossing, and they should be used in any serious backcrossing program if resources are available to the breeder.

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